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C-102

C-102, suspended in a 20 percent solution of polyvinylpyrrolidone, was orally administered to albino rats at levels of 10.0, 31.6, 100.0, 316.0 and 563.0 mg/kg. Due to the limited amount of material, two animals were used at the lowest level and one animal, each, at all higher dosage levels. Two animals served as controls. The initial weight of the animals ranged between 88 and 104 grams. Animals were weighed (Appendix) and examined for symptoms as required (Table II).

No objective symptoms were observed at the lowest level.

Hypoactivity was apparent at all levels above 10.0 mg/kg within 0.25 to 1.0 hour and lasted from about 3 to less than 24 hours. The animals, although hypoactive, were not lethargic and were easily aroused by such external stimuli as sudden movements or noises. During the state of hypoactivity, the animals were observed to wash their faces more than normal. The male at the 316.0 mg/kg level and the female at the 563.0 mg/kg level, chewed excessively during this period.

The female at the 563.0 mg/kg level was hypersensitive to touch after 1.25 hours and remained hypersensitive for about

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24 hours. The animal became belligerent when pressure was applied to its tail.

Slow respiration was observed for as long as 24 hours in all hypoactive animals at the three highest levels.

Ataxia was observed in the female at the 100.0 mg/kg level and in all animals at higher levels. The onset time decreased from 1.75 to 1.25 hours, while the duration increased from 2 to about 24 hours as the dosage was increased from 100.0 to 563.0 mg/kg.

Poor equilibrium was observed at the two highest levels 1.5 to 2.5 hours after dosing and lasted from about 5 to 24 hours.

Staggering was observed in both animals at the two highest levels. The animals walked with awkward gaits 1.5 to 1.75 hours after receiving C-102. When walking, they dragged the hind quarters while the forelimbs were stiffened as in tetany. Although the animals would not allow their limbs to remain in a given position, they stopped in awkward positions with their hindlimbs extended posteriorly or forelimbs extended anteriorly for periods of 10 to 15 seconds.

Both animals at the two highest levels had a low peripheral temperature 2.75 to 3 hours after administration.

No deaths occurred at the above levels.

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